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ESTROGEN AND PROGESTERONE RECEPTOR STATUS IN PRE-MENOPAUSAL AND POST-MENOPAUSAL WOMEN PRESENTING WITH BREAST CANCER

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ABSTRACT

Objective: To determine the status of estrogen and the progesterone receptor among pre-menopausal and post-menopausal women with breast cancer

Background: Women all around the world are most commonly hit by the malignancy named the “breast cancer”. About one fifth of women in the world die due to this as its incidence is continuously increasing. Age, age at menarche, parity, age at menopause are few of the risk factors responsible. Age and menopause are the two factors which affects the overall influence the ER/ PR status. Because of clear evidence that investigation of the estrogen receptor (ER) expression reveals clue for the biologic and prognostic progression of disease.

Materials And Methods: A cross sectional study was conducted at surgical Unit-III, Chandka Medical College Hospital, Larkana, using tissue sample biopsy 120 patients were evaluated to determine the estrogen receptor (ER) status. Statistical package for social sciences (SPSS) version 25 was used to enter and analyze the data.

Results: In this study, the postmenopausal women were 87 (72.5%) while premenopausal were 33 (27.5%) in this study Main outcome of the study was ER/PR status which was found that among postmenopausal women ER and PR were positive in 77% (n =67) & 79.3% (n = 69) respectively while among premenopausal women ER and PR were positive in 97% (n =32) & 69.7% (n = 23) respectively

Conclusion: The results show that nearly three fourths of all breast cancer affected patients had an ER and PR were positive- while majority of these were postmenopausal women. Education & awareness, screening & early detection and proper and quick treatment response to ER/PR positive and all other subtypes of breast cancer are the key to better prognosis.

Keywords: Breast Cancer, Estrogen Progesterone Receptors, Postmenopause.

INTRODUCTION

Women all around the world are most commonly hit by the malignancy named the “breast cancer”. About one fifth of women in the world die due to this as its incidence is continuously increasing [1]. Age, age at menarche, parity, age at menopause are few of the risk factors responsible [2, 3, 6]. More than two centuries have passed since the breast cancer has been identified and it was treated by removing the ovaries of affected women. Though; it was associated with increase in life of patients, but the relative mechanism had not been understood for improved survival. Some thought it is a nervous connection of ovaries with the breast tissues [4, 5, 8], but later on,

estrogen and progesterone receptors -presented in the breast-were evidenced to be the ultimately responsible to ligand-receptor pathway. Age and menopause are the two factors which affects the overall influence the ER/ PR status [10] of such hormone dependant tumours. Studies have found strong correlation between the ER positive rate and the age, histological differentiation, tumor size, and menopause status [9-11]. American Society of Clinical Oncology suggests which is further endorsed by another institute i-e; College of American Pathologists that all the new cases of invasive breast cancer should be tested for diagnosis of breast cancer [12], because of clear evidence that investigation of the estrogen receptor (ER) expression reveals clue for the biologic and prognostic progression of disease [13] Study by Rajan G, et al., found that 47.4% postmenopausal and 56.1% premenopausal women having breast cancer and these women had status of ER positive and 34.7% postmenopausal and 47.7% premenopausal women were PR positive [14]. Very



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different from these statistics, a study by Petricevic J, et al. reported that Estrogen receptors were present in 73.9% & 66.7% postmenopausal and premenopausal women respectively and Progesterone receptors were present in 70.4% & 69.6% premenopausal and postmenopausal with breast cancer respectively [15].

The worldwide practice of assessing the status of E receptors & P receptors for breast cancer is common. Based on the available international data on the ER and PR status and its relationship with the breast cancer and lacking of local data on this very serious issue of women health, there is need for assessment of local population of before and after menopause presenting with breast cancer for ER/ PR expression. The results of the current study discuss the clinical use of ER as a predictive biomarker in order to identify correctly doubted patients of breast cancer.

MATERIAL AND METHODS

A cross sectional study was conducted at surgical Unit-III, Chandka Medical College Hospital, Larkana for a period of six months from 01-07-2018 to 31-12-2018. One hundred and twenty study subjects were included in the study using non probability purposive sampling technique. Married/widowed woman diagnosed case of breast cancer aging from 35 to 75 were part of the study.

Patients included in the study went through evaluation of ER and PR status, to determine the estrogen receptor (ER) status we used the tissue sample biopsy.

Statistical package for social sciences (SPSS) version 25 was used to enter and analyze the data. Descriptive analysis was done using mean & standard deviation (Mean \pm SD), While categorical variables were analyzed using proportions and percentages. Approval for the study was obtained from ethical review committee of the SMBBMU Larkana. All the patients who agreed to participate were asked to complete informed consent forms. The respondents were assured for the privacy of their answers.

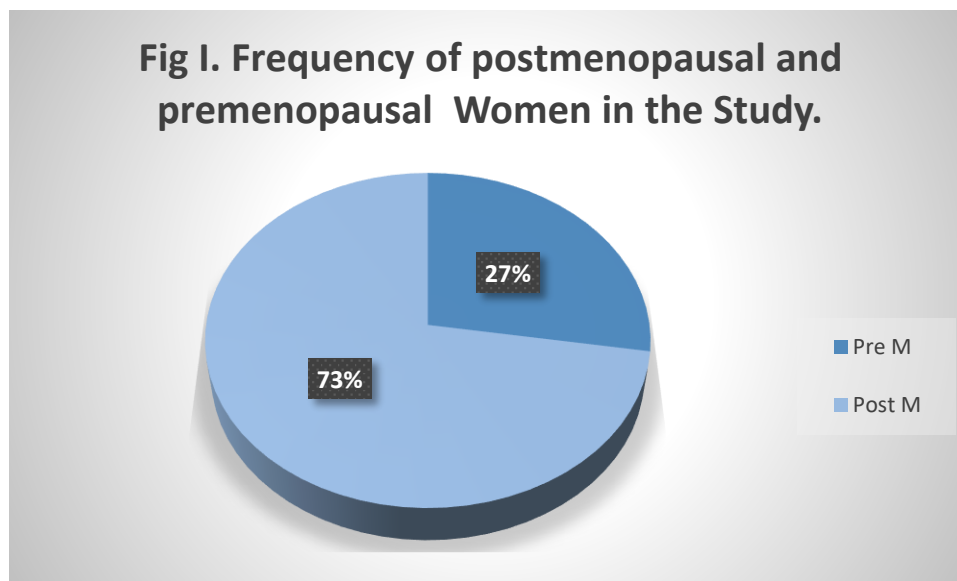
RESULTS

In postmenopausal women the mean \pm SD age was 63.29 ± 8.27 years. The mean \pm SD age while it was 44.61 ± 3.78 years in premenopausal women. The age of menarche was 13.02 ± 1.15 & 13.94 ± 1.17 years in the postmenopausal & premenopausal women respectively. The duration of marriage which was 43.69 ± 8.92 & 24.39 ± 3.88 years respectively in the two groups. Mean and standard deviation of variables like parity, number of children, number of abortions, age at menopause, height, weight, BMI, & duration of oral contraceptive use are detailed in table I.

Table I. Baseline Descriptive Data of all Women in Both Groups

Variable	Group	Min	Max	Mean	S D
Age (years)	Post-M	40	75	63.29	8.27
	Pre-M	39	51	44.61	3.78
Age at menarche	Post-M	11	16	13.02	1.15
	Pre-M	12	16	13.94	1.17
Marriage duration	Post-M	20	58	43.69	8.92
	Pre-M	19	31	24.39	3.88
Parity	Post-M	1	12	4.6	2.52
	Pre-M	1	9	4.64	2.01
Live births	Post-M	1	12	4.83	2.54
	Pre-M	2	10	5.03	2.2
Abortions	Post-M	0	2	0.24	0.59
	Pre-M	0	2	0.33	0.69
Age of menopause	Post-M	39	51	44.98	3.35
	Pre-M				
Weight (Kgs)	Post-M	52	98	70.94	9.43
	Pre-M	54	81	67.39	8.7
Height (m ²)	Post-M	2.23	3.39	2.92	0.25
	Pre-M	2.23	3.31	2.9	0.25
BMI	Post-M	17.88	34.16	24.45	3.6
	Pre-M	17.88	28.74	22.73	3.38
Duration of oral contraceptives use (Years)	Post-M	0	12	2.78	3.56
	Pre-M	0	12	3.67	3.77

In this study, the postmenopausal women were 87 (72.5%) while premenopausal were 33 (27.5%) in this study. Fig I



Main outcome of the study was ER/PR status which was found that among postmenopausal women ER and PR were positive in 77% (n =67) & 79.3% (n = 69) respectively while among premenopausal women ER and PR were positive in 97% (n =32) & 69.7% (n = 23) respectively. Table II

A step ahead it was noted that detection of both of ER was 67 in post Menopause and 32 in premenopausal, while PR was positive in 69 and 23

women in post menopause and premenopausal respectively. Much of premenopausal group women 60.6% (n = 20) had history of using contraceptives while 48.3% (n = 42) postmenopausal women had had used contraceptives. The Breast cancer history in family was found more positively among postmenopausal women 69% (n = 60) than the premenopausal women. Table II

Table II. Frequency Distribution of categorical variables in the Study

Study Variable	Group	Frequency	Percent
Status of Subjects	Post M	87	72.5
	Pre M	33	27.5
ER status Positive	Post-M	67	77
	Pre-M	32	97
PR status Positive	Post-M	69	79.3
	Pre-M	23	69.7
Urban	Post-M	50	41.67
	Pre-M	17	14.17
Rural	Post-M	37	30.83
	Pre-M	16	13.33
Positive family history of breast cancer	Post-M	60	69
	Pre-M	21	63.6
Contraceptive Use	Post-M	42	48.3
	Pre-M	20	60.6

The Chi square test of association is detailed in table III reveals the association between ER/PR status and variables in the study. Stratified analysis revealed non-significantly that frequency of ER/PR detection increased with increasing age among both postmenopausal and premenopausal women - however it was worth noting that rise of frequency was much rapid in premenopausal women. (P value = 0.532 & 0.850 respectively. Rural living women of either group (postmenopausal or premenopausal) women had lesser frequency of ER/PR positive

detection compared to urban living (P value = 0.316 & 0.246 respectively. The family history of breast cancer appeared to be the only finding in the study to have significant association with detection of ER /PR. Table III

Table III. Chi Square Association of Various Factors with Status of ER/PR

Study Variable			Er/Pr Detected		Total	P Value	
			Yes	No			
Family History of Breast cancer	Yes	Post M	37	23	60	0.599	
			61.70%	38.30%	100.00%		
		Pre M	13	8	21		
			61.90%	38.10%	100.00%		
			Post M	13	14	27	0.041
				48.10%	51.90%	100.00%	
Pre M		10	2	12			
		83.30%	16.70%	100.00%			
Menarche age	Up to 12 y	Post M	23	10	33	0.266	
			69.70%	30.30%	100.00%		
		No	Post M	0	4		
			100.00%	0.00%	100.00%		
	13-14 y	Post M	22	23	45	0.094	
			48.90%	51.10%	100.00%		
		Pre M	14	6	20		
			70.00%	30.00%	100.00%		
	≥15 y	Post M	5	4	9	0.681	
			55.60%	44.40%	100.00%		
Pre M		5	4	9			
		55.60%	44.40%	100.00%			
Age category	35-45	Post M	2	2	4	0.532	
			50%	50%	100%		
		Pre M	13	8	21		
			61.90%	38.10%	100%		
	46-55	Post M	5	4	9	0.85	
			55.60%	44.40%	100.00%		
Pre M		10	2	12			
		83.30%	16.70%	100.00%			
Menarche age	Up to 12 years	Post M	23	10	33	0.266	
			69.70%	30.30%	100.00%		
		Pre M	4	0	4		
			100.00%	0.00%	100.00%		
	13-14 years	Pre M	14	23	45	0.094	
			70.00%	51.10%	100.00%		
				6	20		
				30.00%	100.00%		
	15 years and above	Post M	5	4	9	0.681	
			55.60%	44.40%	100.00%		
Pre M		5	4	9			
		55.60%	44.40%	100.00%			
BMI	Underweight	Post M	3	3	6	0.455	
			50.00%	50.00%	100.00%		

		Pre M	6	3	9	
			66.70%	33.30%	100.00%	
	Normal weight	Post M	29	23	52	0.429
			55.80%	44.20%	100.00%	
	Overweight	Pre M	10	6	16	
			62.50%	37.50%	100.00%	
	Post M	15	9	24	0.193	
			62.50%	37.50%	100.00%	
		Pre M	7	1	8	
			87.50%	12.50%	100.00%	
Oral Contraceptive Use	Yes	Post M	23	19	42	0.456
			54.80%	45.20%	100.00%	
	Pre M	12	8	20		
		60.00%	40.00%	100.00%		
No	Post M	27	18	45	0.091	
		60.00%	40.00%	100.00%		
Pre M		11	2	13		
		84.60%	15.40%	100.00%		
Residence	Urban	Post M	30	20	50	0.316
			60.00%	40.00%	100.00%	
	Pre M	12	5	17		
		70.60%	29.40%	100.00%		
Rural	Post M	20	17	37	0.246	
		54.10%	45.90%	100.00%		
Pre M	11	5	16			
		68.80%	31.20%	100.00%		
No. od Abortions	One abortion	Post M	46	34	80	0.195
			57.50%	42.50%	100.00%	
	Pre M	20	9	29		
		69.00%	31.00%	100.00%		
Two abortions	Post M	4	3	7	0.53	
		57.10%	42.90%	100.00%		
Pre M	3	1	4			
		75.00%	25.00%	100.00%		
No. Of Children	Up to 2 children	Post M	8	7	15	0.588
			53.30%	46.70%	100.00%	
	Pre M	2	1	3		
		66.70%	33.30%	100.00%		
	3-5 children	Post M	27	19	46	0.279
			58.70%	41.30%	100.00%	
Pre M	14	6	20			
		70.00%	30.00%	100.00%		
5-10 children	Post M	13	10	23	0.371	
		56.50%	43.50%	100.00%		
Pre M	7	3	10			
		70.00%	30.00%	100.00%		

DISCUSSION

Breast cancer is the number one killer disease of women all over the world. More than twenty percent of deaths in female population are due to malignant breast cancers. The disease affects both the perimenopausal and postmenopausal women, and it is rarely found in women of age younger age i-e; below 25 years [7-11, 17, 21]

Studies and researchers postulate that estrogen is main role player in development of breast cancer as the normal breast tissue contains estrogen (ER) and progesterone (PR) receptors [16, 18, 19, 20]. Family history of breast cancer in women an inverse picture was seen with statistical significance. (P value = 0.599 & 0.041 respectively) .Increasing parity (number of children) (P values = 0.588, 0.279, 0.371, & 0.155 respectively), COC use (P value = 0.456 & 0.091) and rural living (P value = 0.316 & 0.246 respectively) were two protective factors and were associated with lesser frequency of ER/PR positive detection. Study by Russo J, et al., and other few studies have found that pregnancy and parity are two important protective factors [24]. Well defined factor which increasing chances of disease [12, 15, 22, 23, 24] Same was the findings of the current study but with slight variation. It was interesting to note that ER/PR positive detection was on higher side among postmenopausal women in presence of positive family history while in premenopausal.

CONCLUSION

The results show that nearly three fourths of all breast cancer affected patients had an ER and PR were positive- while majority of these were postmenopausal women. Education & awareness, screening & early detection and proper and quick treatment response to ER/PR positive and all other subtypes of breast cancer are the key to better prognosis.

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